

**Amendments to the Claims :**

This listing of claims will replace all prior versions, and listings, of claims in the application.

**Listing of Claims:**

1. (Currently amended) Apparatus for electroporation comprising a wave generator, a biochip containing an array of microelectrodes and a control system that permits to transfer the of a signal to a pre-selected single microelectrode of the biochip.
2. (Currently amended) Apparatus according to claim 1 ~~characterised in that~~ wherein the control system consists of a personal computer equipped with a software program capable of designing various waveform signals and a switching system controlling the wave generator output.
3. (Currently amended) Apparatus according to claims 1 and 2 ~~characterised in that~~ wherein the biochip comprises an array of microelectrodes of a size comparable to a the cell to be electroporated and each of said microelectrodes being driven separately from the others allowing very precise and punctual control of the electroporation process.
4. (Currently amended) A biochip Biochip comprising an array of individually driven microelectrodes (20) comprised on a suitable insulating layer mounted on a solid substrate; means to electrically connect said microelectrodes to a switching system; a cell culture chamber where the cells can be grown and adhere in contact with said array of microelectrodes on a surface formed by said insulating layer containing said array of microelectrodes on said solid substrate.
5. (Currently amended) The biochip Biochip according to claim 4 comprising a semiconductor substrate as the solid substrate covered with an insulating layer (27) comprising said array of individually driven microelectrodes (20) of a size comparable to the cell to be electroporated, and mounting a cell culture chamber (24) with an opening (26) mounted, in turn, on a support (21) made of dielectric

material, said microelectrodes (20) being electrically connected via conductive traces (28) to conductive pads (29) electrically connected, in turn, to a couple of external parallel connectors (22) through wire bonding (23) covered by an the outer portion of the cell culture chamber (24) encircling the opening (26), being said cell culture chamber (24) with the opening (26) mounted over the top of the said semiconductor substrate covered with the an insulating layer (27), both attached on the dielectric support (21).

6. (Currently amended) A biochip Biochip according to claim 5 comprising two further electrodes (25) integrated in the semiconductor substrate covered with the an insulating layer (27), and acting as a ground reference.
7. (Currently amended) A biochip Biochip according to claim 5 wherein the semiconductor substrate covered with the an insulating layer (27) is a silicon substrate covered with an a insulating layer preferentially of SiO<sub>2</sub>.
8. (Currently amended) The biochip Biochip according to claim 5 wherein the these solid substrates are is transparent.
9. (Currently amended) The biochip Biochip according to claim 5 wherein the dielectric support is vetronite, glass or ceramic.
- 10.(Currently amended) The biochip Biochip according to claim 5 wherein the microelectrodes of the array (20) have a size with a surface of at least ten per cent of the total cell membrane and preferably a diameter ranging from 1  $\mu$ m to 50  $\mu$ m.
- 11.(Currently amended) The biochip Biochip according to claims 4 –10 wherein the microelectrodes are of conductive or capacitive type.
- 12.(Currently amended) The microelectrodes Microelectrodes according to claim 11 consisting of conductive microelectrodes obtained over a silicon substrate (31) covered with an a insulating layer preferentially of SiO<sub>2</sub> (32), said microelectrodes having connecting traces wherein said microelectrodes and their connecting traces

(38) being made by a "sandwich" of two titanium nitride, (TiN), layers (33) and an aluminium layer (34), covered with a gold layer (37) on their active surface.

13. (Currently amended) The microelectrodes Microelectrodes according to claim 11 wherein said microelectrodes are realized realised using Metal Oxide Semiconductor, (MOS), technology.

14. (Currently amended) The microelectrodes Microelectrodes according to claim 13 consisting of a silicon p-type substrate (40) in which two n-doped regions, a drain (41) and a source (42), are implanted with conventional microelectronic techniques, the a gate (43) of these electrodes being realized realised in n+ doped polysilicon and is common to all devices in a row, word line, the drain (41) of all devices in a column being connected together by using a metal contact plug and a metal line (44), the source (42) of the transistor being connected via a metal, usually tungsten, plug (46) to a gold layer (47) which acts as the active electrode.

15. (Currently amended) The microelectrodes Microelectrodes according to claim 11 consisting of a capacitive microelectrode obtained with an insulating substrate (60), a metal (61) and a thin insulating layer (64) said microelectrodes being separated by insulating material (62) and covered in non exposed areas by a passivation layer (63).

16. (Currently amended) A method Method of electroporation characterised in that wherein an apparatus according to claims 1 — 3 is used.

17. (Currently amended) The method Method according to claim 16 characterised in that wherein said apparatus performs one or more electroporations to at least a single adhering cell.

18. (Currently amended) The method Method according to claim 17 characterised in that wherein said apparatus comprises a biochip according to claims 4 — 11.

19. (Currently amended) The method Method according to claims 16 — 18 characterised

in that wherein said biochip comprises microelectrodes according to claims 11 12—15.

20. (Currently amended) The method Method according to claims 16 —19 characterised in that wherein the wave generator sends to the electrodes trains of pulses of various amplitude and duration.

21. (Currently amended) The method Method according to claims 16— 20 characterised in that wherein the wave generator sends to the microelectrodes five trains of 25 pulses, {1 ms duration}, repeated at a time interval of 500 ms.

22. (Currently amended) The method Method according to claims 16—20 characterised in that wherein a trains of triangular voltages consisting of 10 pulses are applied to the electrodes the interval between one train and another being of 5 s.

23. (Currently amended) The method Method according to any of preceding claims 16 characterised in that it comprisesing substantially the following steps:

- cultivating cultivate cells since the adhering stage is reached;
- adding add in the culture medium at least one compound to be electroporated in at least one single cell of the said cells;
- selecting selected at least one single cell and at least one microelectrode on which said selected single cell is adherent;
- generating generate at least one electric signal suitable capable to electroporate said at least one single cell with said at least one compound to be electroporated and driving drive said electric signal to the said one microelectrode on which said selected single cell is adherent.

24. (Canceled) Electroporated cells characterised in that they are obtained with method according claims 16—23.

25. (Canceled) Electroporated cells according to claim 24 wherein the electroporated agents are drugs, genetic constructs and proteins.

26. (New) The method Method according to claim 23 wherein the said compounds to be electroporated are drugs, genetic constructs and proteins.

